CVD Risk Factor Disparities in Youth

UNTHSC Health Disparities Conference

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Disclosure

- NHLBI Career Development Award (K23)
- I have no industry disclosures
- No unapproved therapies will be discussed



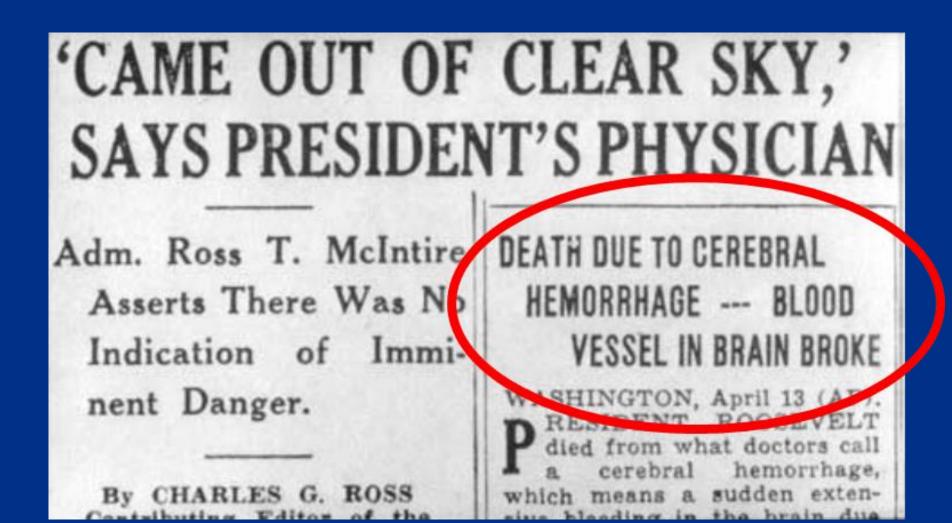
Roadmap

- Pop Quiz- Historical digression
- CVD definition
- CVD developmental progression
- Risk Factors and Subclinical phenotypes
- Pediatric perspectives
- Do disparities drive our understanding?









Messerli. New England Journal of Medicine 1995





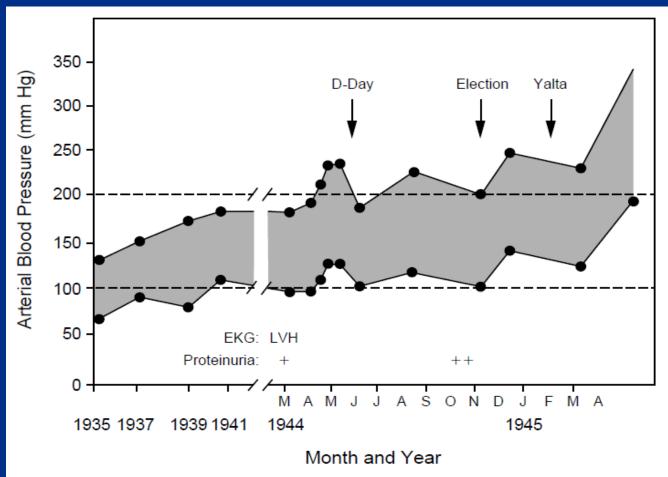


Figure 2. Diastolic and Systolic Arterial Pressure of Franklin D. Roosevelt from 1935 until His Death on April 12, 1945.

EKG denotes electrocardiogram, and LVH left ventricular hypertrophy. Data are from the diary of Dr. Howard G. Bruenn.²

Messerli. New England Journal of Medicine 1995





Definitions

- Cardiovascular disease events
 - Coronary: Heart attack/myocardial infarction, revascularization, coronary disease, angina
 - Cranial: Stroke (ischemic vs. hemorrhagic), transient ischemic attacks
 - Heart failure: hospitalizations or clinical symptoms
 - Other: Sudden death, arterial rupture, arrhythmia, etc.



Pathophysiologic progression

CVD risk factors

Subclinical phenotypes

CVD events

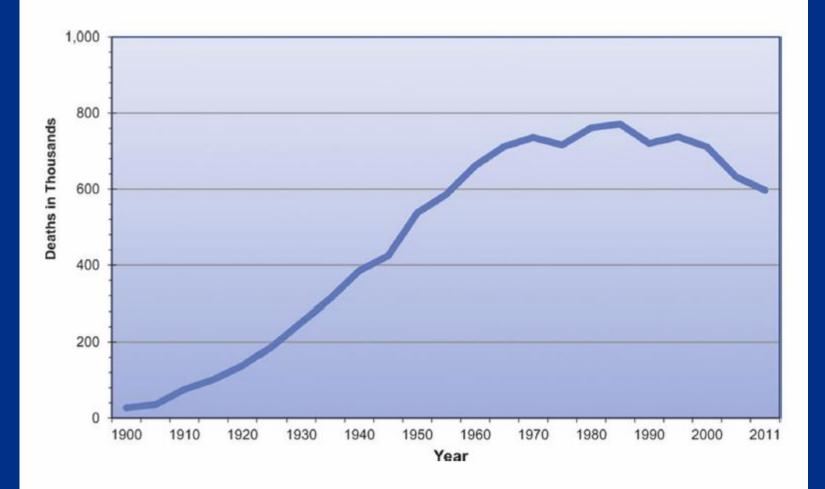


Classic CVD Risk Factors: Framingham Risk Score

- Age
- Sex
- Systolic Blood Pressure
- Total Cholesterol [=HDL+LDL+Triglycerides/5]
- HDL Cholesterol
- Smoking
- Diabetes Mellitus



Deaths attributable to diseases of the heart (United States: 1900-2011).



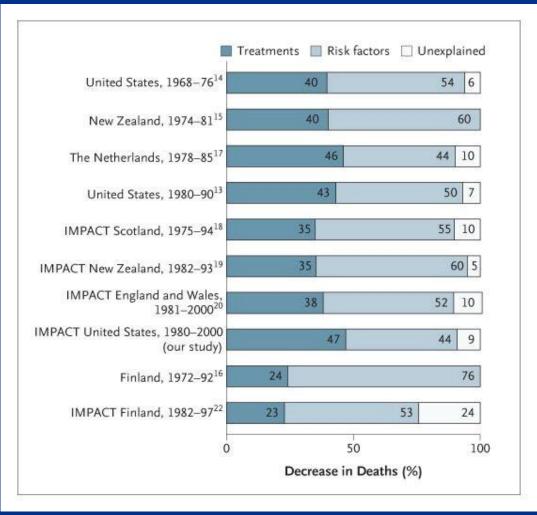


Mozaffarian D et al. Circulation. 2015;131:e29-e322 Copyright © American Heart Association, Inc. All rights reserved.





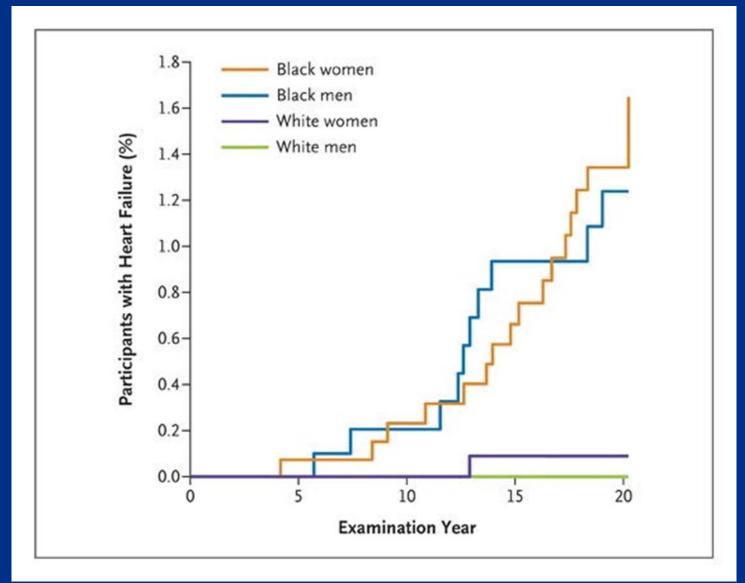
% Decrease in CHD death from 'Prevention' or 'Rescue'



Ford ES et al. N Engl J Med 2007;356:2388-2398.



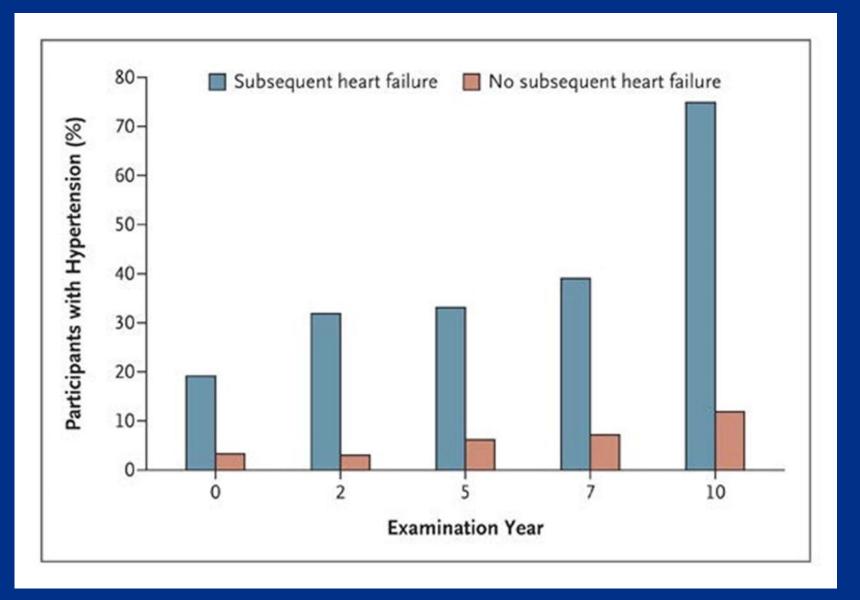




Bibbins-Domingo K et al. N Engl J Med 2009;360:1179-1190.







Bibbins-Domingo K et al. N Engl J Med 2009;360:1179-1190





Subclinical phenotypes

- Vascular
 - Stiffness
 - Thickening
 - Plaque deposition
 - Loss of reactivity
- Cardiac
 - Hypertrophy (Remodelling)
 - Decreased function
 - Metabolic changes

- Brain
 - Vascular
 - Metabolic
 - Structural
- Kidney
 - Proteinuria
 - Structural
 - Vascular
- Others



Implications of left ventricular mass

- Adjusted for age, diastolic BP, Pulse Pressure, HTN treatment, smoking, Diabetes mellitus, obesity, Total:HDL-C ratio, EKG LV hypertrophy
- RR [95%CI] per each additional 50g

- Men
 - CVD 1.49 [1.20-1.85]
 - CVD Death 1.73 [1.19-2.52]
 - All Death 1.49[1.14-1.94]
- Women
 - CVD 1.57 [1.20-2.04]
 - CVD death 2.12 [1.28-3.49]
 - All Death 2.01 [1.44-2.81]

Levy et al. NEJM 1990:322:1561-6



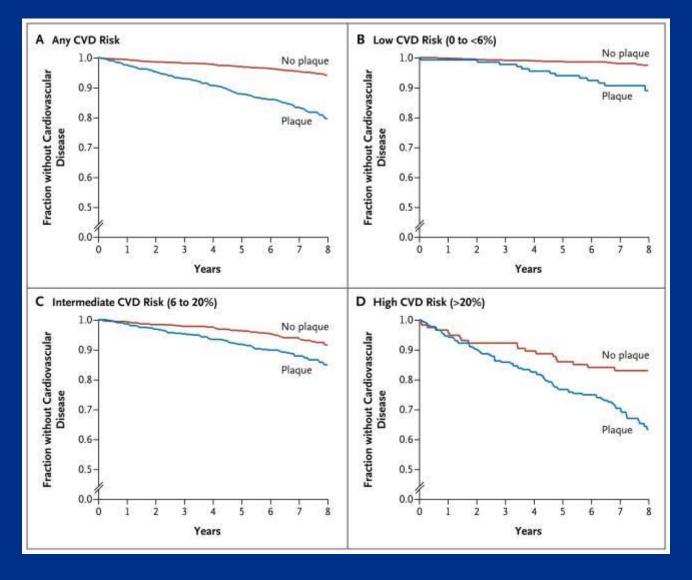


Study o	haracteris	tics	A Total	CV events	βВ	CV n	nortality	:	С	All-cause n	nortality	
Author F	opulation	RR	95% CI	RR (95% CI)	RR	95% CI	RR (95% CI)		RR	95% CI	RR (95% CI)	
Anderson 2009	GEN			1	_				1.15	1.01 - 1.31	-0-	
Blacher 1999	ESRD	1.17	1.06 - 1.30	-0-	1.17	1.06 - 1.30	-0-		1.17	1.07 - 1.27	-0-	
Cruickshank 2002	DM	****	****			****			1.08	1.03 - 1.12	0	
Laurent 2001	HTN	1.09	1.02 - 1.16	-0-	1.09	1.02 - 1.16	□		1.06	1.01 - 1.16	ō	
Meaume 2001	GEN	1.19	1.03 - 1.37	-0-	1.19	1.03 - 1.37	-0-				_	
Pannier 2005	ESRD	1.12	1.03 - 1.22	-0-	1.12	1.03 - 1.22	-0-					
Shoji 2001	ESRD	1.18	1.01 - 1.39		1.18	1.01 - 1.39	-0-		1.15	1.03 - 1.29	-0-	
Shokawa 2005	GEN	1.35	1.13 - 1.61		1.35	1.13 - 1.61			1.28	1.16 - 1.41	-0-	
Sutton-Tyrrell 2005	GEN	1.03	1.01 - 1.06		1.03	1.01 - 1.06			1.05	1.02 - 1.08	h -	
Wang 2010 (men)	GEN	1.21	1.10 - 1.33	-0-	1.21	1.10 - 1.33	-0-		1.19	1.13 - 1.26		
Wang 2010 (women)	GEN	1.30	1.20 - 1.42	-0-	1.30	1.20 - 1.42	-0-		1.25	1.19 - 1.32	1 70	
Willum-Hansen 2006	GEN	1.05	1.01 - 1.08		1.06	1.01 - 1.11					"	
Zoungas 2007	ESRD	1.14	1.07 - 1.22	-0-			(5000)					
Overall		1.14	1.09 - 1.20	•	1.15	1.09 - 1.21	•		1.15	1.09 - 1.21	•	
		0.5		1 2	0.5		1	2	0.5		1	
	Test for heterogeneity: I ² =81.1%, P<0.001 Test for overall effect: Z=5.43, P<0.001						eity: I²=81.3%, P<0.00 ect: Z=4.88, P<0.001	01			eity: I ² =85.5%, P< ect: Z=5.00, P<0.0	

Vlachopoulos et al. Journal of the American College of Cardiology. 2010; 55:1318 - 1327







Polak et al. N Engl J Med. 2011 Jul 21; 365(3): 213-221



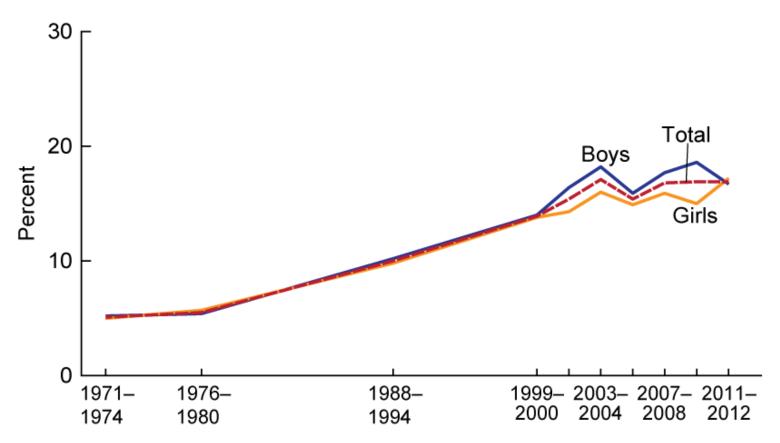


Kids are the future (of CVD prevention)

- Physiologic plasticity
- Developmental advantages
- Fewer political obstacles



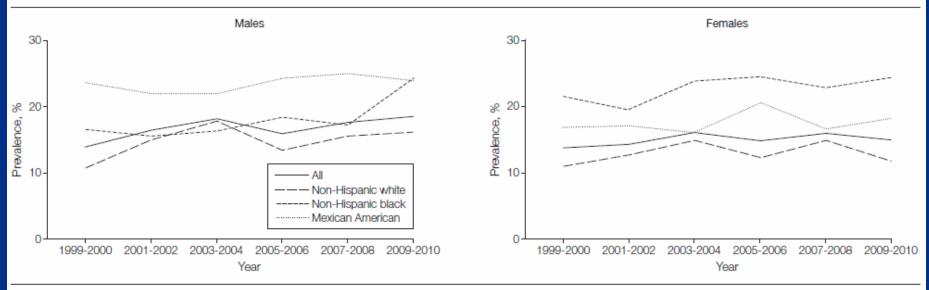
Figure. Trends in obesity among children and adolescents aged 2–19 years, by sex: United States, selected years 1971–1974 through 2011–2012



NOTE: Obesity is body mass index greater than or equal to the sex- and age-specific 95th percentile from the 2000 CDC Growth Charts. SOURCE: CDC/NCHS, National Health and Nutrition Examination Surveys 1971–1974; 1976–1980; 1988–1994; 1999–2000, 2001–2002, 2003–2004, 2005–2006, 2007–2008, 2009–2010, and 2011–2012



Figure 2. Prevalence of Obesity in US Males and Females Aged 2 Through 19 Years



Data are weighted.

488 JAMA, February 1, 2012-Vol 307, No. 5

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	% (95% CI)								
	2-19 y	2-5 y	6-11 y	12-19 y					
Overweight or Obese (BMI for Age ≥85th Percentile of the CDC Growth Charts)									
All race/Hispanic origin groups ^b									
All	31.8 (29.1-34.7)	22.8 (18.7-27.6)	34.2 (30.1-38.5)	34.5 (30.1-39.2)					
Boys	32.0 (29.2-35.0)	23.9 (20.1-28.2)	33.2 (27.7-39.1)	35.1 (29.7-40.9)					
Girls	31.6 (27.2-36.5)	21.7 (14.6-31.0)	35.2 (29.2-41.8)	33.8 (27.9-40.4)					
Non-Hispanic white									
All	28.5 (24.0-33.4)	20.9 (14.4-29.2)	29.4 (21.6-38.7)	31.2 (24.3-39.1)					
Boys	27.8 (22.5-33.8)	21.8 (14.9-30.8)	26.5 (18.2-36.8)	31.5 (21.9-42.9)					
Girls	29.2 (22.7-36.7)	19.9 (10.0-35.7)	32.7 (19.8-48.8)	31.0 (22.7-40.7)					
Non-Hispanic black									
All	35.2 (30.2-40.6)	21.9 (16.7-28.2)	38.1 (30.1-46.8)	39.8 (32.9-47.2)					
Boys	34.4 (30.3-38.7)	22.2 (16.9-28.6)	39.3 (30.5-48.9)	37.3 (30.3-44.9)					
Girls	36.1 (28.7-44.4)	21.6 (14.6-30.8)	36.9 (26.9-48.1)	42.5 (31.9-53.8)					
Non-Hispanic Asian									
All	19.5 (15.7-23.9)	9.0 (4.5-17.3) ^{c,d}	19.9 (16.2-24.3)	24.6 (17.8-32.9)					
Boys	25.1 (18.7-32.8)	8.3 (2.5-24.0) ^{c,d}	24.5 (16.6-34.5)	33.9 (23.2-46.6)					
Girls	13.7 (8.6-21.2)	9.7 (3.3-25.0) ^{c,d}	14.9 (8.9-23.9)	15.0 (7.3-28.3) ^d					
Hispanic									
All	38.9 (36.3-41.6)	29.8 (24.0-36.4)	46.2 (41.5-50.9)	38.1 (31.9-44.8)					
Boys	40.7 (37.3-44.1)	31.4 (23.5-40.5)	48.7 (41.1-56.3)	39.6 (31.3-48.5)					
Girls	37.0 (33.4-40.8)	28.1 (19.7-38.3)	43.6 (37.5-49.8)	36.5 (28.8-45.0)					

Ogden et al. JAMA. 2014;311:806-14.





				% (95%CI)				P Value
Characteristic	1999-2000	2001-2002	2003-2004	2005-2006	2007-2008	2009-2010	2011-2012	for Linear Trend
High TC, ≥200 mg/dL								
Total	10.6 (8.3-13.2)	9.7 (7.1-12.8)	9.8 (7.8-12.2)	9.3 (8.0-10.8)	7.4 (5.6-9.6)	7.0 (5.2-9.2)	7.8 (5.7-10.4)	.006
Boys	9.1 (6.0-13.0)	9.5 (5.3-15.5)	9.1 (6.4-12.4)	10.1 (7.2-13.5)	8.2 (5.4-11.9)	7.8 (5.4-10.9)	6.6 (4.5-9.1)	.14
Girls	12.1 (9.4-15.3)	9.8 (8.1-11.7)	10.7 (7.6-14.5)	8.5 (6.1-11.4)	6.6 (4.5-9.2)	6.1 (4.0-8.8)	9.0 (5.4-13.8)	.007
Low HDL-C, <40 mg/dL								
Total	17.9 (15.0-21.0)	18.7 (16.7-20.8)	11.4 (8.6-14.8)	10.6 (8.6-12.9)	15.6 (13.7-17.7)	13.8 (11.7-16.1)	12.8 (9.8-16.2)	.003
Boys	19.2 (15.7-23.1)	23.0 (19.8-26.4)	15.7 (11.3-20.9)	12.1 (9.0-15.8)	18.3 (15.8-21.1)	16.7 (14.1-19.5)	12.6 (10.3-15.2)	<.001
Girls	16.4 (11.9-21.8)	14.1 (11.8-16.7)	7.0 (4.9-9.6)	9.0 (7.0-11.4)	12.8 (9.8-16.2)	10.8 (7.2-15.5)	12.9 (8.3-18.9)	.30
High non-HDL-C, ≥145 mg/dL								
Total	13.6 (11.3-16.2)	14.6 (12.3-17.1)	10.6 (8.2-13.3)	10.6 (9.2-12.2)	10.0 (7.5-12.9)	8.5 (6.2-11.4)	8.4 (5.9-11.5)	<.001
Boys	13.1 (10.3-16.3)	15.6 (12.2-19.5)	10.4 (6.8-15.1)	10.3 (7.8-13.2)	11.5 (7.9-15.9)	9.0 (6.6-11.9)	7.5 (5.6-9.9)	<.001
Girls	14.2 (11.3-17.5)	13.5 (10.9-16.6)	10.8 (7.7-14.5)	10.9 (8.2-14.2)	8.3 (5.8-11.5)	8.0 (4.9-12.2)	9.2 (5.6-14.1)	.001





		% (95% CI)				
Characteristic	Participants, No.	TC ≥200 mg/dL	HDL-C <40 mg/dL ^b	Non-HDL-C ≥145 mg/dL ^c	High TC, Low HDL-C, or High Non-HDL-C ^d	
Total	1482	7.8 (5.7-10.4)	12.8 (9.8-16.2)	8.4 (5.9-11.5)	20.2 (16.3-24.6)	
Sex						
Boyse	757	6.6 (4.5-9.1)	12.6 (10.3-15.2)	7.5 (5.6-9.9)	19.3 (16.2-22.8)	
Girls	725	9.0 (5.4-13.8)	12.9 (8.3-18.9)	9.2 (5.6-14.1)	21.0 (14.8-28.5)	
Age, y						
8-12 ^e	785	7.0 (4.4-10.4)	10.5 (6.9-15.1)	6.9 (4.2-10.7)	18.1 (12.5-24.9)	
13-17	697	8.5 (5.2-12.9)	14.7 (11.7-18.1)	9.6 (6.1-14.2)	22.0 (18.0-26.6)	
Race/Hispanic origin ^f						
Non-Hispanic white ^e	346	7.0 (3.7-11.9)	13.9 (9.5-19.3)	7.7 (3.8-13.5)	19.7 (13.7-27.1)	
Non-Hispanic black	433	10.0 (8.3-11.8)	5.6 (3.5-8.5)	9.1 (7.5-10.9)	17.6 (15.1-20.4)	
Non-Hispanic Asian	176	7.5 (3.7-13.1)	9.2 (6.0-13.3)	7.0 (2.6-14.6) ^g	16.6 (11.2-23.3)	
Hispanic	452	8.6 (6.0-11.8)	15.7 (10.9-21.7)	10.4 (7.8-13.5)	24.2 (19.0-29.9)	

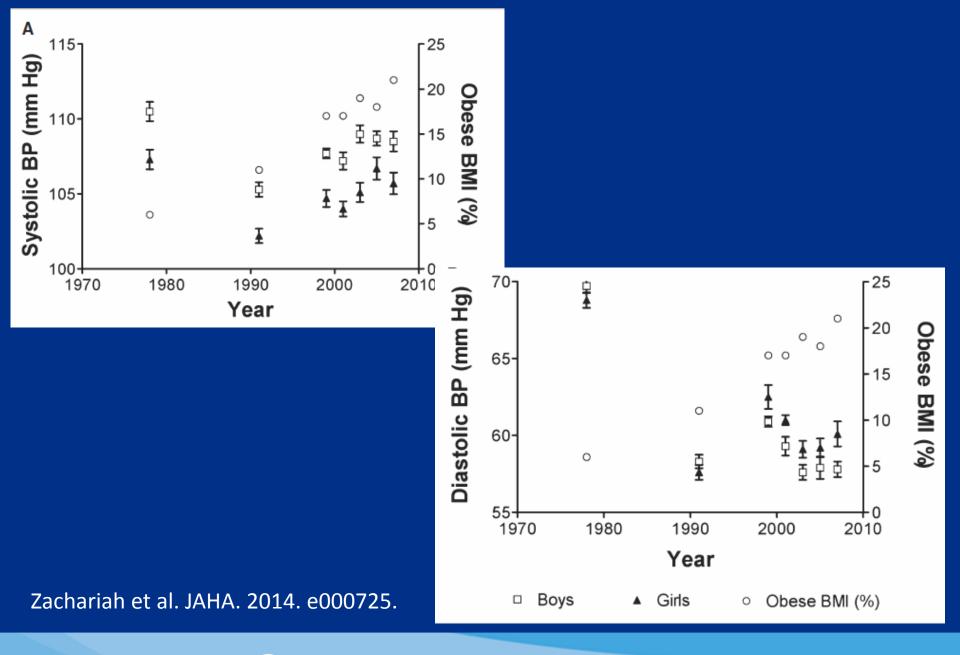




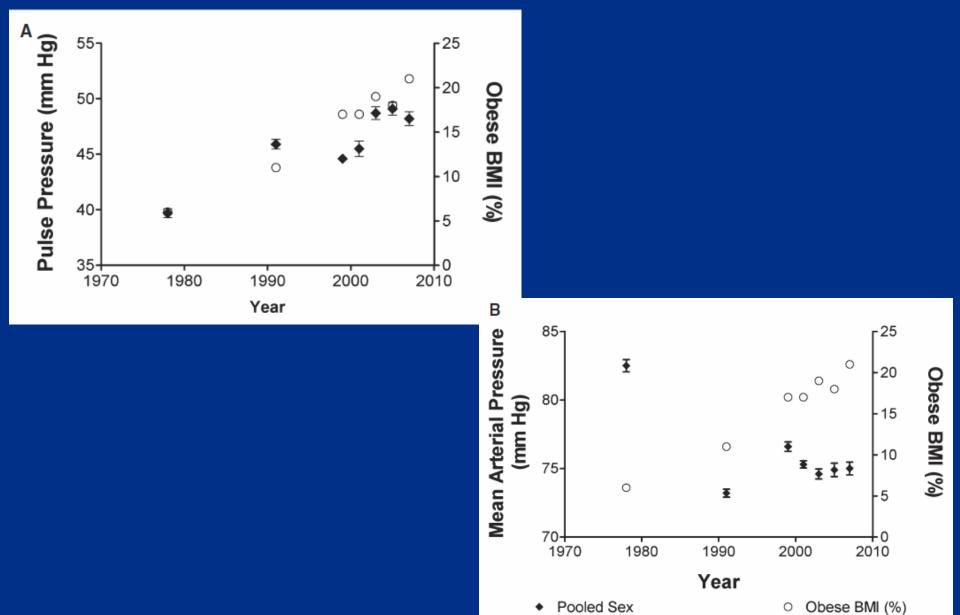
High BP								
Total	3.0 (2.0-4.3)	2.7 (1.7-4.1)	3.1 (1.9-4.8)	2.8 (1.5-4.8)	2.6 (1.8-3.5)	1.7 (1.2-2.5)	1.6 (1.0-2.4)	.003
Boys	3.3 (2.0-5.1)	3.2 (2.1-4.6)	3.1 (1.6-5.4)	2.0 (1.0-3.5)	3.2 (2.0-4.8)	1.6 (0.9-2.7)	1.8 (0.6-4.1) ^b	.03
Girls	2.7 (1.0-6.0) ^c	2.3 (1.2-4.0)	3.1 (1.5-5.7)	3.6 (1.8-6.6)	2.0 (0.7-4.2) ^c	1.9 (1.3-2.7)	1.4 (0.8-2.1)	.11
Borderline high BP								
Total	7.6 (5.8-9.8)	10.0 (8.3-11.9)	9.1 (7.4-11.1)	10.3 (7.7-13.4)	10.1 (8.3-12.2)	7.2 (5.4-9.3)	9.4 (7.2-11.9)	.90
Boys	10.0 (7.8-12.7)	14.1 (11.2-17.5)	13.2 (10.2-16.8)	13.9 (10.0-18.7)	12.5 (8.8-17.0)	10.5 (7.5-14.1)	13.7 (9.5-18.8)	.74
Girls	5.1 (2.9-8.3)	5.7 (4.1-7.7)	4.9 (3.6-6.5)	6.5 (4.2-9.5)	7.7 (5.7-10.0)	3.7 (2.2-5.9)	5.4 (3.0-9.0)	.95
High or borderline high BP								
Total	10.6 (8.4-13.1)	12.7 (10.6-15.1)	12.2 (9.8-15.0)	13.1 (9.4-17.5)	12.7 (10.6-15.0)	8.9 (7.2-10.9)	11.0 (8.8-13.4)	.26
Boys	13.3 (10.7-16.3)	17.3 (14.4-20.5)	16.3 (12.5-20.6)	15.9 (11.3-21.5)	15.6 (11.8-20.0)	12.1 (9.2-15.5)	15.4 (11.0-20.9)	.61
Girls	7.9 (4.3-12.9)	8.0 (5.8-10.7)	8.0 (5.4-11.4)	10.2 (6.7-14.6)	9.7 (7.0-12.8)	5.6 (4.3-7.3)	6.8 (4.0-10.6)	.42













			% (95% CI)	
Characteristic	Participants, No.b	High BP	Borderline High BPc	Either High or Borderline High BP ^d
Total	1665	1.6 (1.0-2.4)	9.4 (7.2-11.9)	11.0 (8.8-13.4)
Sex				
Boyse	842	1.8 (0.6-4.1) ^f	13.7 (9.5-18.8)	15.4 (11.0-20.9)
Girls	823	1.4 (0.8-2.1)	5.4 (3.0-9.0)	6.8 (4.0-10.6)
Age, y				
8-12 ^e	904	1.9 (1.1-3.0)	4.7 (2.7-7.4)	6.5 (4.5-9.1)
13-17	761	1.3 (0.5-2.8) ^g	13.7 (10.3-17.7)	15.0 (11.2-19.4)
Race/Hispanic origin ^h				
Non-Hispanic white ^e	388	1.1 (0.5-1.9)	8.3 (5.5-12.0)	9.4 (6.7-12.7)
Non-Hispanic black	483	1.9 (0.6-4.3) ^f	13.5 (10.5-17.0)	15.3 (12.5-18.6)
Hispanic	502	2.4 (0.7-5.6) ^f	9.1 (4.3-16.4)	11.5 (6.3-18.7)
Non-Hispanic Asian	203	1.7 (0.5-4.2) ^f	6.9 (3.3-12.4)	8.5 (3.8-16.0) ^g





Table 3. Suggested Revised Schema for Staging of Ambulatory BP Levels in Children

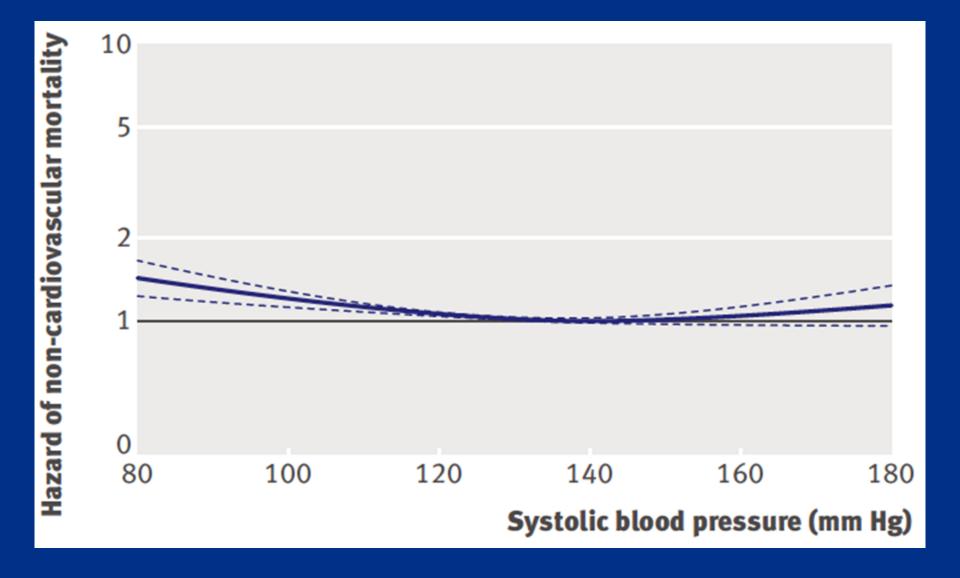
Oleanities kinn	Office DD+	Mean Ambulatory	CDD DDD 0/ +C
Classification	Office BP*	SBP or DBP†‡	SBP or DBP Load, %‡§
Normal BP	<90th %tile	<95th %tile	<25
White coat hypertension	≥95th %tile	<95th %tile	<25
Prehypertension	\geq 90th %tile or >120/80 mm Hg	<95th %tile	≥25
Masked hypertension	<95th %tile	>95th %tile	≥25
Ambulatory hypertension	>95th %tile	>95th %tile	25–50
Severe ambulatory hypertension (at risk for end-organ damage)	>95th %tile	>95th %tile	>50

marker for renal deterioration. Racial differences also have been demonstrated in nocturnal dipping, with a difference in the relationship between body size and BP contributing to the elevated nighttime pressures seen in African American as compared with white youth.⁹²

Flynn et al. Hypertension 2014. 63:1116-35. Urbina et al. Hypertension 2008. 52:433-51.



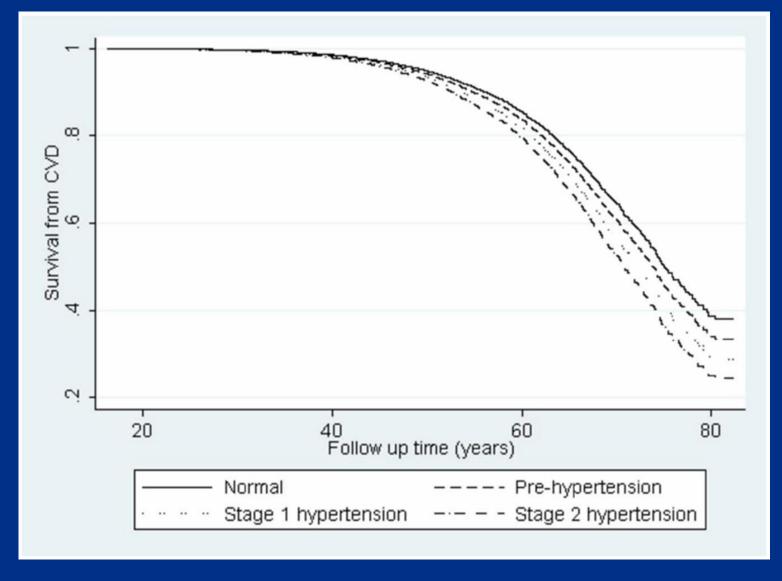




Sundstrom et al. British Medical Journal 2011.







Gray et al. Journal of the American College of Cardiology 2011.





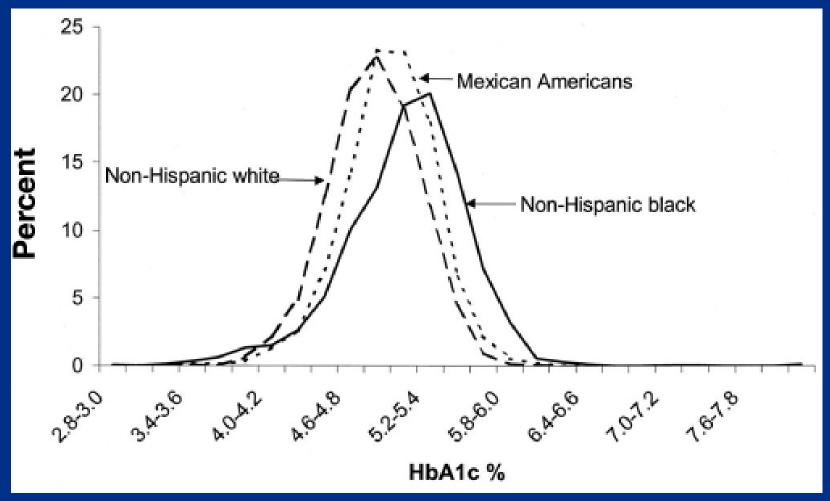
	All T2DM ^a			Dia	Diagnosed T2DM ^a			Undiagnosed T2DM ^a		
	No. of Persons ^b	% ^c	95% CI ^c	No. of Persons ^b	% ^c	95% CI ^c	No. of Persons ^b	% ^c	95% CI ^c	
Total population	119,224	0.36	0.20, 0.67	78,613	0.24	0.11, 0.51	40,611	0.12	0.05, 0.31	
Sex										
Male	74,523	0.44	0.19, 1.02	44,087	0.26	0.08, 0.82	30,436	0.18	0.06, 0.55	
Female	44,701	0.28	0.13, 0.60	34,526	0.22	0.09, 0.53	10,175	0.06	0.01, 0.29	
Race/ethnicity										
Non-Hispanic white	56,171	0.28	0.10, 0.80	44,783	0.22	0.07, 0.72	11,388	0.06	0.01, 0.40	
Non-Hispanic black	18,904	0.40	0.14, 1.14	12,819	0.27	0.07, 1.07	6,084	0.13	0.03, 0.50	
Mexican-American	27,386	0.73	0.40, 1.40	17,737	0.48	0.22, 1.04	9,649	0.26	0.09, 0.75	
Other Hispanic	0			0			0			
Other ^d	16,764	0.77	0.17, 3.40	3,274	0.15	0.01, 2.04	13,490	0.62	0.11, 3.45	

Demmer et al. Am J Epi 2013.178:1106-13.





HgbA1c in NHANES ages 5–24



Herman. J Diabetes Sci Technol. 2009; 3: 656–660.





Left ventricular Hypertrophy: Child to Young Adult

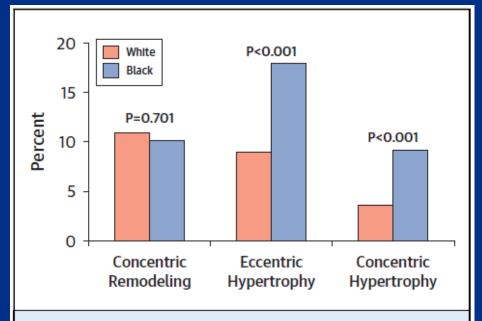


FIGURE 2 Prevalence of LV Remodeling Patterns, by Race

Concentric remodeling, eccentric hypertrophy, and concentric hypertrophy were defined by cutoffs of left ventricular (LV) mass index (46.7 g/m $^{2.7}$ in women and 49.2 g/m $^{2.7}$ in men) and relative wall thickness (0.42).

Lai et al. JACC Imaging 2014; 64:1580-7.



TABLE 2	Effects of LVMI a	nd LVH on BMI and SB	P
(Linear an	nd Logistic Regress	sion Analyses)	

	t Variable	
Independent Variable	LVMI*	LVH†
Childhood (Model I‡)		
BMI	0.26§ (0.20-0.32)	1.65§ (1.39-1.97)
SBP	0.08 (0.01-0.14)	1.27 (1.04-1.54)
Adulthood age	0.16§	1.09§
Sex	-0.10§	1.14
Race	0.16§	2.58§
Adulthood (Model II¶)		
BMI	0.42§ (0.37-0.48)	2.53§ (2.06-3.09)
SBP	0.16§ (0.10-0.21)	1.56§ (1.28-1.90)
Adulthood age	0.14§	1.10§
Sex	-0.11§	1.12
Race	0.17§	2.97§
Total AUC (Model III#)		
BMI	0.41§ (0.36-0.47)	2.42§ (1.98-2.95)
SBP	0.14§ (0.09-0.20)	1.47§ (1.20-1.80)
Adulthood age	0.15§	1.12§
Sex	-0.10§	1.16
Race	0.17§	2.96§
Incremental AUC (Model IV**)		
BMI	0.33§ (0.28-0.39)	2.09§ (1.72-2.53)
SBP	0.10§ (0.05-0.16)	1.43§ (1.19-1.72)
Adulthood age	0.17§	1.13§
Sex	-0.10§	1.13
Race	0.17§	2.80§

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Take Home

- CVD events are declining
- Risk factor management is a key driver
- Abnormal CVD Risk Factor prevalence in youth is high, but may be stable or declining
- Racial Disparities are present in
 - Abnormal CVD risk factor prevalence
 - Subclinical atherosclerotic precursors
- Disparities may lead to CVD event differences



Thank you for your attention.

Questions?



